## STN SEARCH SUMMARY 10/826,729

```
=> d his
      (FILE 'HOME' ENTERED AT 17:18:17 ON 29 MAR 2005)
     FILE 'REGISTRY' ENTERED AT 17:18:30 ON 29 MAR 2005
                  E JUGLONE/CN
L1
                1 S E3
                  E FREDERICAMYCIN-A/CN
                  E FREDERICAMYCIN/CN
L2
                1 S E5
                  E PIN-1/CN
L3
                1 S E7
     FILE 'CAPLUS' ENTERED AT 17:20:49 ON 29 MAR 2005
L4
            1236 S L1 OR L2
L5
               27 S L4 AND PIN?
               30 S L4 AND (PIN? OR PPIASE? OR ISOMERASE)
L6
L7
               30 S L4 AND (PIN? OR PPIASE? OR ISOMERASE OR PARVULIN)
                3 S L7 AND PARVULIN
L8
               15 S L4 AND ((CELL W PROLIFERATION) OR (ABNORMAL W CELL W GROWTH)
L10
L11
              397 S (PIN1 OR PIN-1) OR (PIN W 1)
                                                                                          (CANCOR))
L12
           31466 S (PIN1 OR PIN-1) OR (PIN W 1) OR PIN
L13
              169 S L12 AND ISOMERASE
              172 S L12 AND (ISOMERASE OR PEPTIDYL OR PARVULIN)
L14
L15
               98 S L14 AND (MODULAT? OR INHIBIT? OR ACTIVAT?)
                1 S L15 AND COVALENT
L16
=> e juglone/cn
                      JUGLOMYCIN Z/CN
               1
            JUGLONE ACETATE/CN

JUGLONE ACETATE/CN

JUGLONE BENZYL ETHER/CN

JUGLONE GLUCOSIDE/CN

JUGLONE HYDROXYLASE/CN

JUGLONE METHYL ETHER/CN

JUGLONE OXIDOREDUCTASE/CN

JUGLONE REDUCTASE/CN

JUGLONE, 2,3,7-TRIMETHOWN

JUGLONE
E2
E3
E4
E5
E6
E7
E8
E9
E10
E11
                      JUGLONE, 2,3,7-TRIMETHOXY-/CN
E12
                      JUGLONE, 2,3,7-TRIMETHOXY-, ACETATE/CN
=> s e3; d
               1 JUGLONE/CN
L1
     ANSWER 1 OF 1 REGISTRY COPYRIGHT 2005 ACS on STN
RN
      481-39-0 REGISTRY
ED
     Entered STN: 16 Nov 1984
     1,4-Naphthalenedione, 5-hydroxy- (9CI) (CA INDEX NAME)
```

OTHER CA INDEX NAMES:

Juglone (6CI)

CN

CN 1,4-Naphthoquinone, 5-hydroxy- (8CI)

```
OTHER NAMES:
CN
     1,4-Dihydro-1,4-dioxo-5-hydroxynaphthalene
CN
     5-Hydroxy-1, 4-naphthalenedione
CN
     5-Hydroxy-1, 4-naphthoquinone
CN
     5-Hydroxynaphthoquinone
CN
     8-Hydroxy-1, 4-naphthoguinone
CN
     Akhnot
     C.I. 75500
CN
     C.I. Natural Brown 7
CN
CN
     Juglon
CN
     NSC 153189
CN
     NSC 34266
CN
     NSC 622948
CN
     Nucin
CN
    , Regianin
CN
     Walnut Extract
FS
     3D CONCORD
MF
     C10 H6 O3
CI
     COM
                  AGRICOLA, ANABSTR, AQUIRE, BEILSTEIN*, BIOBUSINESS, BIOSIS,
LÇ
     STN Files:
       BIOTECHNO, CA, CABA, CANCERLIT, CAOLD, CAPLUS, CASREACT, CEN, CHEMCATS,
       CHEMINFORMRX, CHEMLIST, CIN, CSCHEM, DDFU, DRUGU, EMBASE, GMELIN*,
       HODOC*, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK*, NAPRALERT,
       NIOSHTIC, PHAR, PROMT, RTECS*, SPECINFO, SYNTHLINE, TOXCENTER, USPAT2,
       USPATFULL
         (*File contains numerically searchable property data)
                     DSL**, EINECS**
     Other Sources:
         (**Enter CHEMLIST File for up-to-date regulatory information)
=> e fredericamycin/cn
             1
                   FRECIDE/CN
E2
             1
                   FRED/CN
E3
             0 --> FREDERICAMYCIN/CN
E4
             1
                   FREDERICAMYCIN <SYM65>-CYCLODEXTRIN 1:2 COMPLEX/CN
E5
             1
                  FREDERICAMYCIN A/CN
E6
             1
                 FREDERICAMYCIN A TETROL/CN
E7
             1
                 FREDERICAMYCIN B/CN
E8
             1
                 FREDERICAMYCIN C/CN
                 FREDERICAMYCIN TETROL/CN
E9
             1
                 FREDERICAMYCIN TETROL TRIPOTASSIUM SALT/CN
E10
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                  FREDERICAMYCIN-3-CARBOXALDEHYDE/CN
E11
             1
E12
                 FREDERICON A/CN
=> s e5;d
             1 "FREDERICAMYCIN A"/CN
     ANSWER 1 OF 1 REGISTRY COPYRIGHT 2005 ACS on STN
L2
RN
     80455-68-1 REGISTRY
ED
     Entered STN: 16 Nov 1984
     Spiro[2H-benz[f]indene-2,8'-[8H]cyclopent[g]isoquinoline]-1,1',3,5,8(2'H)-
CN
     pentone, 6',7'-dihydro-4,9,9'-trihydroxy-6-methoxy-3'-[(1E,3E)-1,3-
     pentadienyl]-, (2S)- (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
```

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CN
      Spiro[2H-benz[f]indene-2,8'-[8H]cyclopent[g]isoquinoline]-1,1',3,5,8(2'H)-
      pentone, 6',7'-dihydro-4,9,9'-trihydroxy-6-methoxy-3'-(1,3-pentadienyl)-,
      [S-(E,E)]-
 OTHER NAMES:
 CN
      Fredericamycin A
      NSC 305263
 CN
      STEREOSEARCH
 FS
 MF
      C30 H21 N O9
 CI
      COM
 LC
      STN Files:
                   ADISINSIGHT, ANABSTR, BEILSTEIN*, BIOBUSINESS, BIOSIS,
        BIOTECHNO, CA, CANCERLIT, CAPLUS, CASREACT, CEN, CHEMINFORMRX, CIN,
        DDFU, DRUGU, EMBASE, IPA, MEDLINE, MRCK*, NAPRALERT, PHAR, PROMT,
        SYNTHLINE, TOXCENTER, USPATFULL
          (*File contains numerically searchable property data)
 Absolute stereochemistry.
 Double bond geometry as shown.
 => s e7; d
              1 "PIN1 (HUMAN CELL LINE HELA GENE PIN1)"/CN
      ANSWER 1 OF 1 REGISTRY COPYRIGHT 2005 ACS on STN
 L3
      479476-95-4 REGISTRY
 RN
 ED
      Entered STN: 17 Jan 2003
      Pin1 (human cell line HeLa gene PIN1) (9CI) (CA INDEX NAME)
 CN
 OTHER NAMES:
 CN
      1997: PN: WO03095618 TABLE: 1 claimed protein
 CN
      2064: PN: WO03091391 FIGURE: 20 unclaimed protein
 CN
      2841: PN: WO03038130 FIGURE: 3 claimed protein
      387: PN: WO2004038376 TABLE: 5 unclaimed protein
 CN
 CN
      GenBank AAC50492
 CN
      GenBank AAC50492 (Translated from: GenBank U49070)
 FS
      PROTEIN SEQUENCE
      Unspecified
 MF
 CI
      MAN
- SR
      GenBank
 LC
      STN Files: CA, CAPLUS, TOXCENTER, USPATFULL
 => file caplus
 => d 18 2-3
      ANSWER 2 OF 3 CAPLUS COPYRIGHT 2005 ACS on STN
 L8
      2005:54187 CAPLUS
 AN
      142:149781
 DN
 ΤI
      Use of chaperonin PPIase for enhancement of poorly expressed
      proteins and immobilization of the proteins for drug screening
 IN
      Ideno, Akira; Furuya, Masahiro
 PA
      Sekisui Chemical Co., Ltd., Japan
      Jpn. Kokai Tokkyo Koho, 41 pp.
 SO
      CODEN: JKXXAF
```

LA Japanese FAN.CNT 1 PΙ

Patent

DT

PATENT NO. KIND DATE APPLICATION NO. DATE ------------------------JP 2005013067 A2 20050120 JP 2003-181394 20030625 PRAI JP 2003-181394 20030625

ANSWER 3 OF 3 CAPLUS COPYRIGHT 2005 ACS on STN L8

ΑN 1998:276736 CAPLUS

DN 129:51272

ΤI Selective inactivation of parvulin-like peptidyl-prolyl cis/trans isomerases by juglone

ΑU Hennig, Lars; Christner, Claudia; Kipping, Marc; Schelbert, Birte; Rucknagel, Karl Peter; Grabley, Susanne; Kullertz, Gerd; Fischer, Gunter

Enzymology of Protein Folding, Max-Planck Research Unit, Halle/Saale, CS D-06120, Germany

SO Biochemistry (1998), 37(17), 5953-5960 CODEN: BICHAW; ISSN: 0006-2960

PB American Chemical Society

DTJournal

LA English

RE.CNT 44 THERE ARE 44 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

## => d 110 10-15

L10 ANSWER 10 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN

ΑN 1994:499158 CAPLUS

DN 121:99158

The inhibitory action of juglone on tumor cell multiplication TI

ΑU Zhang, Yeping; Yang, Zhibo; Jing, Yongkui; Xu, Shaohui

CS Dep. Pharmacol., Shenyang Coll. Pharm., Shenyang, Peop. Rep. China

SO Shenyang Yaoxueyuan Xuebao (1993), 10(4), 271-4 CODEN: SYXUE3; ISSN: 1000-1727

DTJournal

LA Chinese

L10 ANSWER 11 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN

AN 1992:152406 CAPLUS

DN 116:152406

TIPreparation of LH-RH analogs as hormone-dependent neoplasm inhibitors

IN Schally, Andrew Victor; Janaky, Tamas; Juhasz, Atilla; Bajusz, Sandor

PΑ

SO Eur. Pat. Appl., 26 pp.

CODEN: EPXXDW

DT Patent

LA English

FAN.CNT 4

KIND APPLICATION NO. PATENT NO. DATE DATE \_\_\_\_ ----------PΙ EP 450461 19911009 EP 1991-104730 A2 19910326 EP 450461 A3 19920311 В1 19950906 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE ES 2076393 T3 19951101 ES 1991-104730 19910326 CA 2039908 AA19911007 CA 1991-2039908

	AU 9174106	A1	19911010	AU 1991-74106	19910405
	AU 638319	B2	19930624		
	ни 57235	A2	19911128	HU 1991-1116	19910405
	JP 04224600	A2	19920813	JP 1991-72936	19910405
	ZA 9104552	Α	19920624	ZA 1991-4552	
	WO 9222322	A1	19921223	WO 1991-US4264	19910614
	W: FI, KR, NO				
	NO 9304541	Α	19940207	NO 1993-4541	19931210
PRAI	US 1990-505517	Α	19900406		
		Α	19910614		
os	MARPAT 116:152406				
L10	ANSWER 12 OF 15 CA	APLUS (	COPYRIGHT 20	05 ACS on STN	
AN	1988:221606 CAPLUS	S			
DN	108:221606				
ΤI				olyl derivatives, pro-	
		lations	containing	them, and their use ${\tt a}$	s anticancer
	agents				
IN			Jones, Paul	. Spencer; Cooper, Mar	tin Edward
PA	Glaxo Group Ltd.,				
SO	Ger. Offen., 22 pp	•			
	CODEN: GWXXBX				
DT	Patent				
LA	German				
FAN.	CNT 1				
	PATENT NO.		DATE	APPLICATION NO.	DATE
ΡI	DE 3725185	A1	19880204		
	NO 8702963		19880201		
	NL 8701768	Α			
	DK 8703921	A	19880130		
	FI 8703289	A	19880130		19870728
	711 0776000	- 1	1000000	*** 4000 0000	4 0 0 5 0 5 0 0

P AU 8776203 19880204 AU 1987-76203 A1 19870728 AU 604731 В2 19910103 CN 87105778 Α 19880224 CN 1987-105778 19870728 CN 1015059 В 19911211 GB 2195636 Α1 19880413 GB 1987-17864 19870728 GB 2195636 В2 19900530 JP 63093783 Α2 19880425 JP 1987-186790 19870728 HU 1987-3460 HU 46011 A2 19880928 19870728 HU 201936 В 19910128 SE 1987-2986 SE 8702986 Α 19890129 19870728 ES 2007666 Α6 19890701 ES 1987-2210 19870728 US 4851399 Α 19890725 US 1987-78716 19870728 CH 672489 Α 19891130 CH 1987-2885 19870728 BE 1002110 Α4 19900710 BE 1987-839 19870728 AT 8701909 Α 19910415 AT 1987-1909 19870728 SU 1676445 A3 19910907 SU 1987-4203501 19870728 FR 2602233 A1 19880205 FR 1987-10763 19870729 FR 2602233 В1 19900817 ZA 8705592 Α 19880831 ZA 1987-5592 19870729 AT 8702563 Α 19911015 AT 1987-2563 19871008 JP 02000187 A2 19900105 JP 1989-33893 19890215 PRAI GB 1986-18398 19860729 GB 1987-10608 Α 19870505

OS CASREACT 108:221606; MARPAT 108:221606

```
L10 ANSWER 13 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN
```

AN 1987:49879 CAPLUS

DN 106:49879

TI Fredericamycin A derivatives

IN Hasegawa, Hiroshi; Yokoi, Koichi; Narita, Masa; Asaoka, Takemitsu; Kukita, Kenichi; Ishizeki, Seiji; Nakajima, Toshiaki

PA S. S. Pharmaceutical Co., Ltd., Japan

SO Jpn. Kokai Tokkyo Koho, 6 pp.

CODEN: JKXXAF

DT Patent

LA Japanese

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE		
ΡI	JP 61044867	A2	19860304	JP 1984-166283	19840808		
	JP 03004548	B4	19910123				
PRAI	JP 1984-166283		19840808				
os	CASREACT 106:49879						

L10 ANSWER 14 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN

AN 1986:497256 CAPLUS

DN 105:97256

TI Fredericamycin A derivatives

IN Hasegawa, Hiroshi; Yokoi, Koichi; Narita, Masa; Asaoka, Takemitsu; Kukita, Kenichi; Ishizeki, Seiji; Nakajima, Toshiaki

PA S. S. Pharmaceutical Co., Ltd., Japan

SO Jpn. Kokai Tokkyo Koho, 9 pp.

CODEN: JKXXAF

DT Patent

LA Japanese

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE			
ΡI	JP 61044868	A2	19860304	JP 1984-166683	19840809			
	JP 03031193	B4	19910502					
PRAI	JP 1984-166683		19840809					

L10 ANSWER 15 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN

AN 1968:28389 CAPLUS

DN 68:28389

TI Mitotic abnormalities produced by juglone in Ehrlich ascites tumor cells

AU Okada, Tadashi A.; Roberts, Eugene; Brodie, Arnold F.

CS City of Hope Med. Center, Duarte, CA, USA

SO Proceedings of the Society for Experimental Biology and Medicine (1967), 126(2), 583-8

CODEN: PSEBAA; ISSN: 0037-9727

DT Journal

LA English

# => d 110 abs

L10 ANSWER 1 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN

AB The invention provides a method for treating a Pin1-associated state in a subject including administering to a subject an effective amount of a fredericamycin A compound such that the Pin1-associated state is treated. In another aspect, the invention includes the above-described method, wherein

the Pinl-associated state is a cyclin D1 elevated state, neoplastic transformation, and/or tumor growth. In an embodiment, the invention provides the above-described methods, wherein the Pinl-associated state is colon cancer, breast cancer, a sarcoma, a malignant lymphoma, and/or esophageal cancer. The invention also provides a method for treating cyclin D1 overexpression in a subject including administering to a subject an effective amount of a combination of a fredericamycin A compound and a hyperplastic inhibitory agent such that the cyclin D1 overexpression is treated.

#### => d 110 10 abs

- L10 ANSWER 10 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN
- AB The IC50 values of juglone for inhibiting the growth of various lines of cultured mouse cancer cells were: 13.8 <SYM109>g/mL (HeLa cell), 9.8 <SYM109>g/mL (P388 cell), 7.1 <SYM109>g/mL (P388/ADR cell) and 11.6 <SYM109>g/mL (S180
  - cell). Juglone's inhibition of S180 cells was cytocidal and concentration-dependent in nature.

## => d 110 5-10

- L10 ANSWER 5 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN
- AN 2001:257994 CAPLUS
- DN 134:266570
- TI Preparation of luteinizing hormone releasing hormone analogs having a cytotoxic moiety
- IN Janaky, Tamas; Juhasz, Attila; Bajusz, Sandor; Schally, Andrew V.
- PA The Administrators of the Tulane Educational Fund, USA
- SO U.S., 15 pp., Cont.-in-part of U.S. Ser. No. 505,517, abandoned. CODEN: USXXAM
- DT Patent
- LA English
- FAN. CNT 4

ъ.	TIV. CIVI 3						
	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE		
Ρ	I US 6214969	B1	20010410	US 1993-8186	19930125		
	NO 9304541	Α	19940207	NO 1993-4541	19931210		
Ρ	RAI US 1988-260994	B2	19881021				
	US 1989-404667	B2	19890907				
	US 1990-505517	B2	19900406				
	WO 1991-US4264	Α	19910614				
0	S MARPAT 134:266570						

- RE.CNT 10 THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L10 ANSWER 6 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN
- AN 2000:535357 CAPLUS
- DN 133:144904
- TI Caspase cascade-based methods for identifying therapeutically effective antineoplastic agents, compounds so identified, and pharmaceutical compositions
- IN Weber, Eckard; Tseng, Ben Y.; Drewe, John; Cai, Sui Xiong
- PA Cytovia, Inc., USA
- SO PCT Int. Appl., 87 pp.

DT Patent LA English FAN.CNT 2 PATENT NO. KIND DATE APPLICATION NO. DATE --------------PΙ WO 2000045165 20000803 A1 WO 2000-US2329 20000201 W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG EP 1151295 A1 20011107 EP 2000-907081 20000201 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO PRAI US 1999-118102P Ρ 19990201 US 1999-454595 Α 19991207 WO 2000-US2329 W 20000201 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT ANSWER 7 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN AN 1997:126257 CAPLUS DN 126:233190 Antitumor promoting effects of naphthoquinone derivatives on short term ΤI Epstein-Barr early antigen activation assay and in mouse skin carcinogenesis ΑU Kapadia, Govind J.; Balasubramanian, Venkataraman; Tokuda, Harukuni; Konoshima, Takao; Takasaki, Midori; Koyama, Junko; Tagahaya, Kiyoshi; Nishino, Hoyoku CS Department of Pharmaceutical Sciences, College of Pharmacy and Pharmaceutical Sciences, Howard University, Washington, D.C. 20059, USA SO Cancer Letters (Shannon, Ireland) (1997), 113(1,2), 47-53 CODEN: CALEDQ; ISSN: 0304-3835 PB Elsevier DT Journal LA English L10 ANSWER 8 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN ΑN 1996:257730 CAPLUS DN 125:10466 TТ Further model studies related to fredericamycin A: analogs in which ring C is expanded to six atoms, and an examination of the diastereoselectivity of radical spirocyclization ΑU Clive, Derrick L. J.; Kong, Xianglong; Paul, Christine Chua Chem. Dep., Univ. Alberta, Edmonton, AB, T6G 2G2, Can. CS SO Tetrahedron (1996), 52(17), 6085-116 CODEN: TETRAB; ISSN: 0040-4020 PΒ Elsevier DTJournal



LΑ

English

L10 ANSWER 9 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN

CODEN: PIXXD2

```
AN
     1995:678592 CAPLUS
DN
     123:74386
     Preliminary study of the effect of selected Chinese natural drugs on human
ΤI
     ovarian cancer cells
ΑU
     Yu, Jing Jie; Reed, Eddie
CS
     National Cancer Institute, National Institutes Health, Bethesda, MD,
     20892. USA
SO
     Oncology Reports (1995), 2(4), 571-5
     CODEN: OCRPEW; ISSN: 1021-335X
PB
     Oncology Reports
DT
     Journal
     English
LA
L10
     ANSWER 10 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN
ΑN
     1994:499158 CAPLUS
DN
     121:99158
ΤI
     The inhibitory action of juglone on tumor cell multiplication
ΑU
     Zhang, Yeping; Yang, Zhibo; Jing, Yongkui; Xu, Shaohui
CS
     Dep. Pharmacol., Shenyang Coll. Pharm., Shenyang, Peop. Rep. China
     Shenyang Yaoxueyuan Xuebao (1993), 10(4), 271-4
SO
     CODEN: SYXUE3; ISSN: 1000-1727
DT
     Journal
LA
     Chinese
=> d 110 3-4
L10
     ANSWER 3 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN
ΑN
     2003:777766 CAPLUS
DN
     139:292095
ΤI
     Preparation of fredericamycin derivatives for use in treating
     cancer
ΙN
     Abel, Ulrich; Simon, Werner
     Bioleads GmbH, Germany; Biofrontera Discovery GmbH
PΑ
SO
     PCT Int. Appl., 116 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     German
FAN.CNT 1
     PATENT NO.
                         KIND
                                DATE
                                            APPLICATION NO.
                                                                    DATE
                         ----
     WO 2003080582
PΙ
                          Α2
                                20031002
                                            WO 2003-EP2922
                                                                    20030320
     WO 2003080582
                          A3
                                20041209
             AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
             CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
             GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
             LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM,
             PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT,
             TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
             KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,
             FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR,
             BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
     DE 10248451
                          A1
                                20031009
                                            DE 2002-10248451
                                                                    20021017
     CA 2480468
                          AA
                                20031002
                                            CA 2003-2480468
                                                                    20030320
     EP 1503988
                          A2
                                20050209
                                            EP 2003-714862
                                                                    20030320
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R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,

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IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
PRAI DE 2002-10213580
                         Α
                               20020326
     DE 2002-10248451
                         Α
                               20021017
    WO 2003-EP2922
                         W
                               20030320
OS
    MARPAT 139:292095
L10 ANSWER 4 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN
ΑN
    2002:594670 CAPLUS
DN
    137:150218
TΙ
    Methods of inhibiting Pinl-associated states using a fredericamycin A
    compound
IN
    Ping, Lu Kun; Fischer, Gunter
PΑ
    Pintex Pharmaceuticals, Iceland
SO
    PCT Int. Appl., 67 pp.
    CODEN: PIXXD2
DT
    Patent
LA
    English
FAN.CNT 1
    PATENT NO.
                       KIND
                               DATE
                                          APPLICATION NO.
                                                                  DATE
                        ----
                               -----
                                           ------
                                                                  _____
PΙ
    WO 2002060436
                         A2
                               20020808
                                           WO 2001-US50597
                                                                  20011221
    WO 2002060436
                         А3
                               20030123
                         C1
    WO 2002060436
                               20030424
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
            GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
            PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
            UA, UG, US, UZ, VN, YU, ZA, ZM, ZW
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
            KG, KZ, MD, RU, TJ, TM, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB,
            GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA,
            GN, GQ, GW, ML, MR, NE, SN, TD, TG
    CA 2432981
                         AA
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           AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
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                        Т2
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PRAI US 2000-257412P
                         Р
                               20001222
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20011221

=> d 110 11-12 abs

OS

WO 2001-US50597

MARPAT 137:150218

L10 ANSWER 11 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN GI

W

- AB Title peptides X-R1-R2-R3-Ser-R5-R6(Q)-Leu-Arg-Pro-R10-NH2 (I; R1 = pyroglutamic acid residue, 3-(2-naphthyl)-D-alanine residue; R2 = His, 4-chloro-D-phenylalanine residue; R3 = Trp, D-Trp, 3-(3-pyridyl)-D-alanine residue; R5 = Tyr, Arg; R6 = D-Lys, D-Orn; R10 = Gly, D-Ala; X = H, C2-5 alkanoyl; Q = cytotoxic moiety Q4, AQ3, B(Q1)2, B(AQ2)2; A = NH(CH2)nCO, OC(CH2) nCO where n = 2,6; B = HNCH2(CH2) mCH(NH) n(CH2) nCO where m,n = 0, 1; A and B are bound to R6 via CO and CO moiety of A is bonded to an amino group on B for B(AQ2)2; Q1 = D- or L-4-[bis(2-chloroethyl)amino]phenylalanine residue, cyclopropylcarbonyl, aziridine-2-carbonyl; etc.; Q2 = Q1, doxorubicin residue, etc.; Q3 = Q2, methotrexoyl, etc.; Q4 - Q1, methotrexoyl] were prepared as LH-RH analogs useful as hormone-dependent neoplasm inhibitors. Thus, title LH-RH analog II was prepared via coupling of [D-Lys]6LH-RH (preparation via standard solid phase method using a benzhydrylamine resin, Boc-Gly-OH, and the appropriate protected amino acids given) and anthraquinone 2-methylhemiglutarate (preparation from 2-(hydroxymethyl)anthraquinone and qlutaric anhydride qiven). II at 10 <SYM109>q/mL in vitro gave 71% inhibition of 3H-thymidine incorporation into DNA in MCF-7 human breast cancer cells.
- L10 ANSWER 12 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN GI
- The title derivs. I [R1, R2 = H, OH, OP(O)(OH)(OR4); R4 = H, alkyl (un)substituted by OH, alkoxy, cyclic ether, cycloalkyl, R4 = alkenyl, cycloalkyl, aryl, aralkyl, aroylalkyl; <SYM179>1 of R1 and R2 = OP(O)(OH)(OR4); R3 = H, halo, Me] and their salts, useful as anticancer agents (no data), were prepared by 5 methods. A suspension of I (R1 = R3 = H, R2 = ICH2CO2) in THF was treated with H2O and HCl and the mixture stirred and refluxed 18 h to give I (R1 = R3 = H, R2 = OH) which, in THF, was added to NaH in THF and the whole treated with ClP(O)(OPr)2 in THF to give I [R1 = R3 = H, R2 = OP(O)(OPr)2]. This reacted with NaI in refluxing MeCOEt to give I [R1 = R3 = H, R2 = OP(O)(OPr)(ONa)] (II). A dry powder for injection comprised a weight II equivalent to 100 mg acid, 8.8 mg tri-Na citrate, and 0.2 mg citric acid per vial.

=> d 110 13-14 abs

L10 ANSWER 13 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN GI

- \* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY AVAILABLE VIA OFFLINE PRINT \*
- AB Stable fredericamycin A derivs. I (R = H, C1-4 alkyl; R1 = C1-4 alkyl), useful as neoplasm inhibitors, were prepared Thus, fredericyamin A (II) was reduced over 10% Pd/C in THF at room temperature for 10 h, then stirred with Ac2O for 1 h to give 80% III. III was heated with MeI and Ag2O in Me2CO for 1 h to give 56.3% I (R = R1 = Me), whose i.p. administration prolonged the lives of mice transplanted with Ehrlich cancer cells (5

+ 106 cells/animal) in a dose dependent manner. A saline solution of III was more stable than that of II.

L10 ANSWER 14 OF 15 CAPLUS COPYRIGHT 2005 ACS on STN GI

- \* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY AVAILABLE VIA OFFLINE PRINT \*
- AB Stable fredericamycin A derivs. I and II (R = H, EtO2C, acyl; R1 = alkyl), useful as neoplasm inhibitors and bactericides, were prepared Thus, fredericamycin A (III, R = H)(IV) was treated with EtO2CCl in pyridine at 0° under stirring to give 83.% III (R = EtO2C), which was then treated with MeI and Ag2O in anhydrous dioxane at 75-80° under stirring to give 56.3% II (R = EtO2C, R1 = Me), whose i.p. administration prolonged the lives of mice transplanted with Ehrlich cancer cells (5 + 106 cells/animal) in a dose dependent manner. The title compds. also showed stronger antibacterial activities against Saccharomyces ruxii and Piricularia oryzae than IV in vitro.

## => d 116

L16 ANSWER 1 OF 1 CAPLUS COPYRIGHT 2005 ACS on STN

AN 2004:934473 CAPLUS

DN 141:388653

TI Methods of treating pinl associated disorders by covalent modification ofactive site residues

IN Tibbitts, Thomas

PA Pintex Pharmaceuticals, Inc., USA

SO PCT Int. Appl., 41 pp. CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

	PATENT NO.						KIND DATE			APPL	ICAT		DATE						
ΡI	WO 20	WO 2004094601		A2 20041104				,	 WO 2	 004-	 US11	20040416							
	W	: AE,	AG,	AL,	AM,	AT,	AU,	ΑZ,	BA,	BB,	ВG,	BR,	BW,	BY,	ΒZ,	CA,	CH,		
		CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,		
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	R'	W: BW,	GH,	GM,	ΚE,	LS,	MW,	ΜZ,	SD,	SL,	SZ,	ΤZ,	UG,	ZM,	ZW,	AM,	ΑZ,		
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		ES,	FI,	FR,	GB,	GR,	HU,	ΙE,	IT,	LU,	MC,	NL,	PL,	PT,	RO,	SE,	SI,		
		SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,		
		TD,	TG																
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PRAI	RAI US 2003-463810P					20030417													

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ΤI
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     -prolyl cis/trans isomerases by juglone
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     Rucknagel, Karl Peter; Grabley, Susanne; Kullertz, Gerd; Fischer, Gunter
CS
     Enzymology of Protein Folding, Max-Planck Research Unit, Halle/Saale,
     D-06120, Germany
SO
     Biochemistry (1998), 37(17), 5953-5960
     CODEN: BICHAW; ISSN: 0006-2960
PΒ
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DT
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RE.CNT 44
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     The mitotic peptidyl-prolyl isomerase, Pin1,
     interacts with Cdc25 and Plx1
ΑU
     Crenshaw, Donna G.; Yang, Jing; Means, Anthony R.; Kornbluth, Sally
CS
     Department of Pharmacology and Cancer Biology, Duke University Medical
     Center, Durham, NC, 27710, USA
SO
     EMBO Journal (1998), 17(5), 1315-1327
     CODEN: EMJODG; ISSN: 0261-4189
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              ALL CITATIONS AVAILABLE IN THE RE FORMAT
    ANSWER 97 OF 98 CAPLUS COPYRIGHT 2005 ACS on STN
L15
     1998:203534 CAPLUS
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ΤI
     Pin1 binds and regulates mitosis-specific phosphoproteins
     Shen, Minhui; Stukenberg, P. Todd; Kirschner, Marc W.; Lu, Kun Ping
ΑU
     Cancer Biology Program, Div. Hematology/Oncology, Dep. Med., Beth Israel
CS
     Deaconess Med. Center and Div. on Aging, Harvard Med. Sch., Boston, MA,
     02215, USA
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ΤI
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IN
     Hunter, Tony; Lu, Kun Ping
PΑ
     Salk Institute for Biological Studies, USA
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SO

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CODEN: PIXXD2

DT Patent LA English FAN.CNT 1

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## => d 115 abs 96

L15 ANSWER 96 OF 98 CAPLUS COPYRIGHT 2005 ACS on STN AB The cis/trans peptidyl-prolyl isomerase, Pin1

, is a regulator of mitosis that is well conserved from yeast to man. Here, we demonstrate that depletion of Pinl-binding proteins from Xenopus egg exts. results in hyperphosphorylation and inactivation of the key mitotic regulator, Cdc2/cyclin B. We show biochem. that this phenotype is a consequence of Pinl interaction with critical upstream regulators of Cdc2/cyclin B, including the Cdc2-directed phosphatase, Cdc25, and its known regulator, Plx1. Although Pinl could interact with Plx1 during interphase and mitosis, only the phosphorylated, mitotically active form of Cdc25 was able to bind Pinl, an event we have recapitulated using in vitro phosphorylated Cdc25. Taken together, these data suggest that Pinl may modulate cell cycle control through interaction with Cdc25 and its activator, Plx1.